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(54) **APPARATUS AND METHOD FOR QUANTIFICATION OF THE DESYNCHRONIZATION BETWEEN THE CLOCKS OF TWO HBC ACTIVE IMPLANTS**

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(58) **Field of Classification Search**
USPC 375/224, 219; 607/60
See application file for complete search history.

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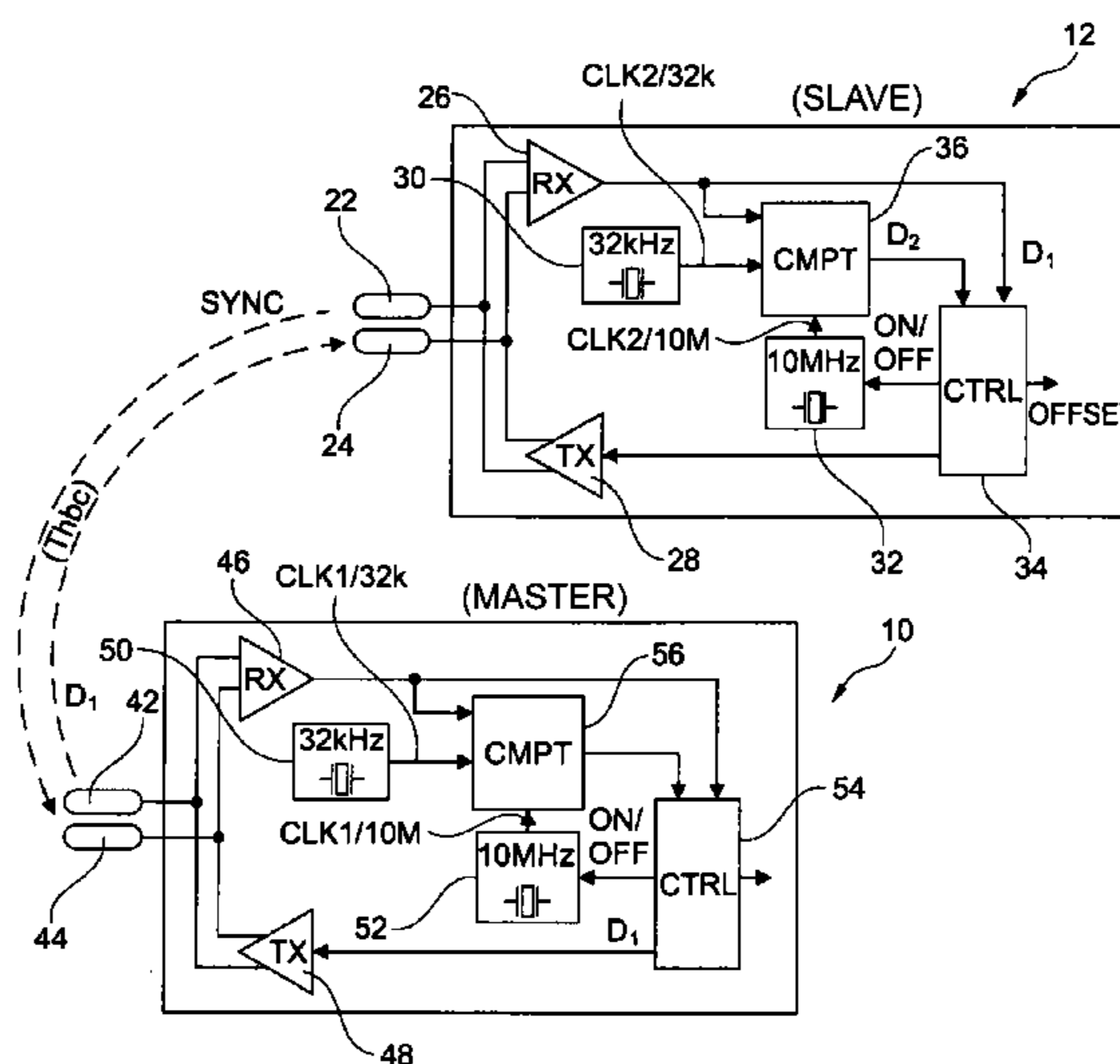
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(57) **ABSTRACT**

A method for quantification of the desynchronization between the clocks of two medical devices communicating wirelessly, for example, by HBC signals. The devices are separately clocked by slow clocks (CLK1/32k, CLK2/32k) and include selectively activated fast clocks (CLK1/10M, CLK2/10M). The method comprises: a) on a predetermined transition (T₁) of a slow clock, transmission by one device of a synchronization query signal (SYNC) to the other device, b) counting of the pulses of the activated fast clock to detect a predetermined transition (T₃) of the first slow clock, then c) transmitting from the other device to the first device a response signal (D₁) and d) upon reception of the response signal, computing a temporal shift (OFFSET) according to the result (D₁, D₂) of the counting of the pulses of the fast clock. Two fast clocks, one on each device, also can be used.

31 Claims, 3 Drawing Sheets



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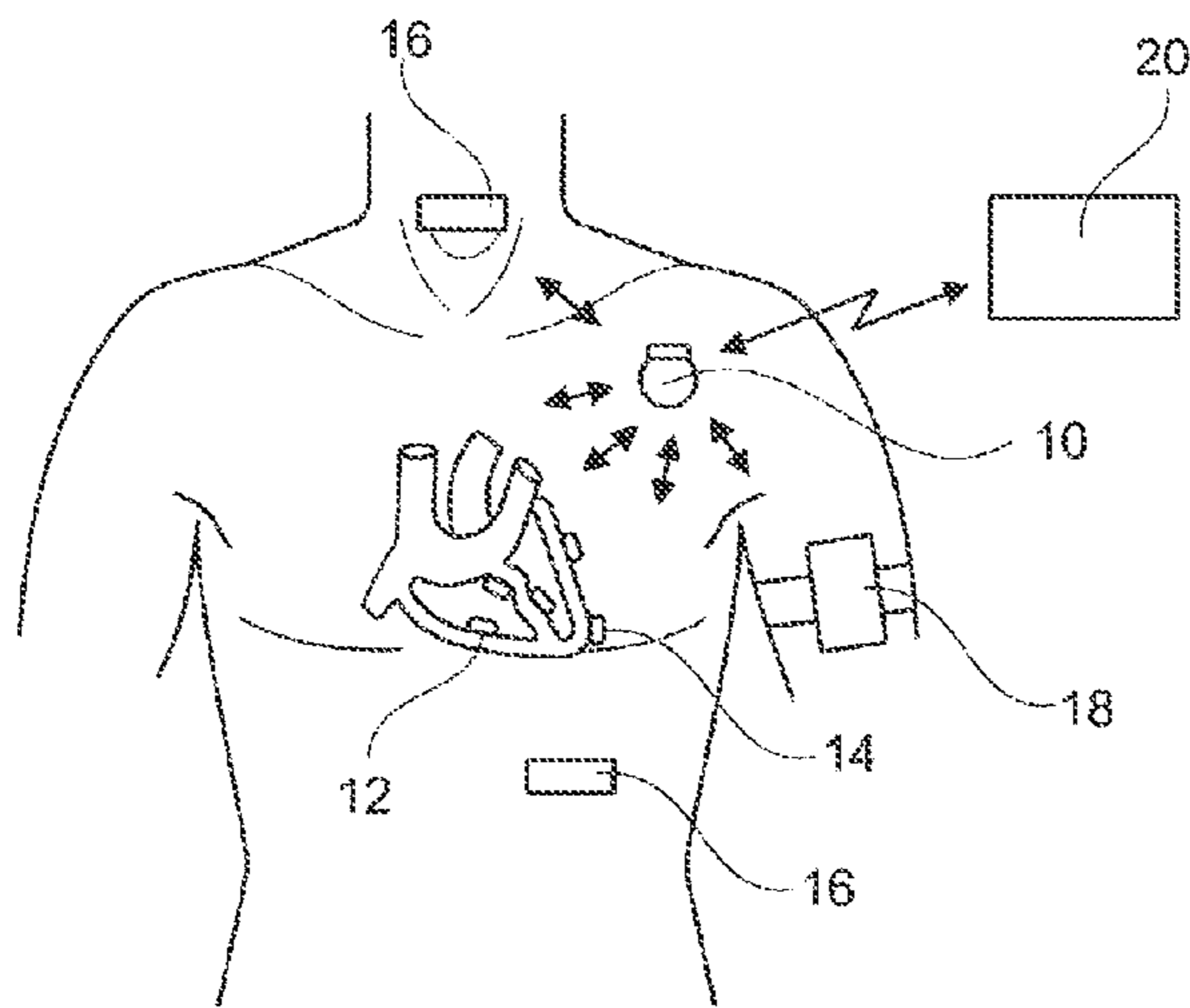


FIG. 1

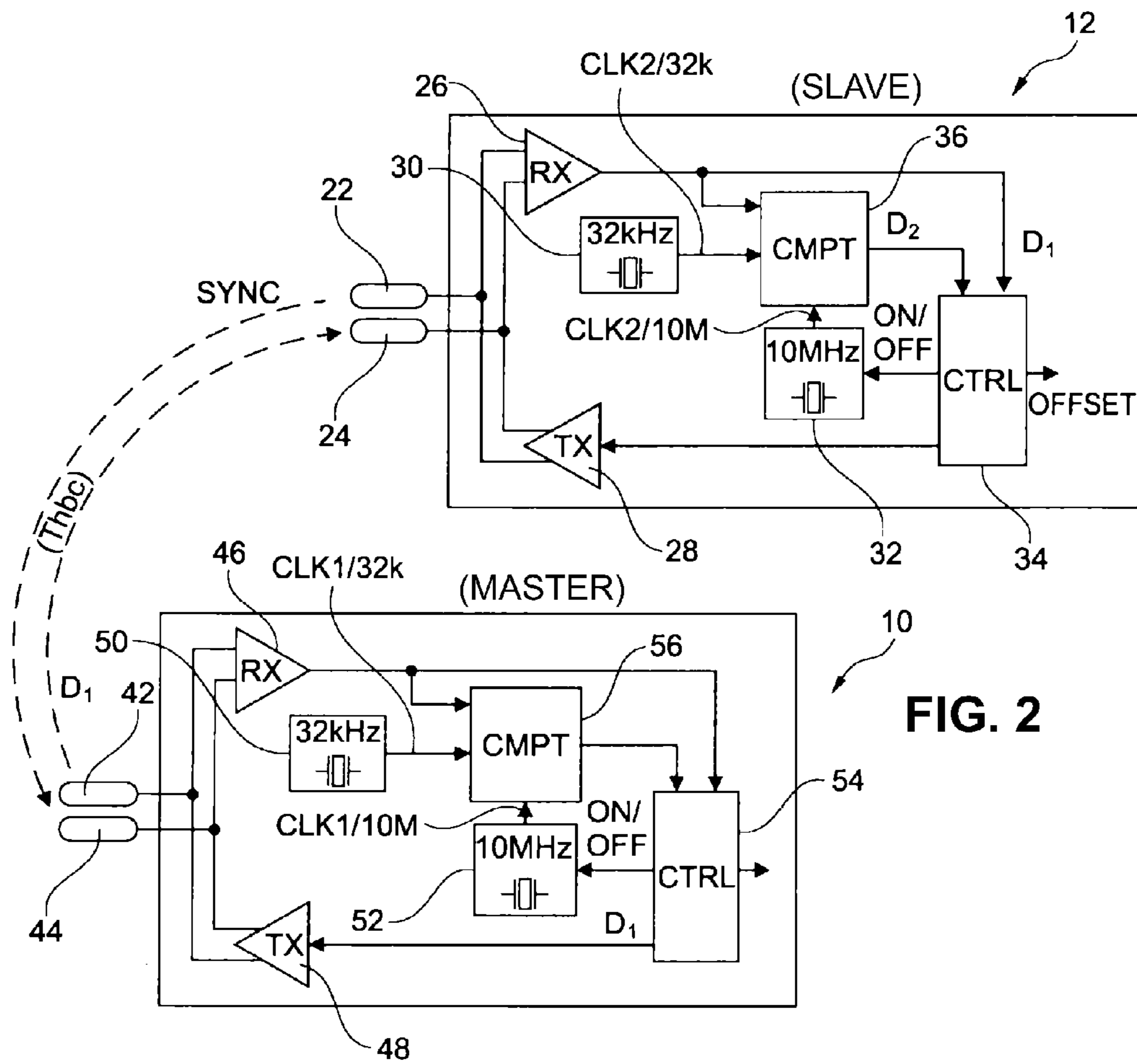


FIG. 2

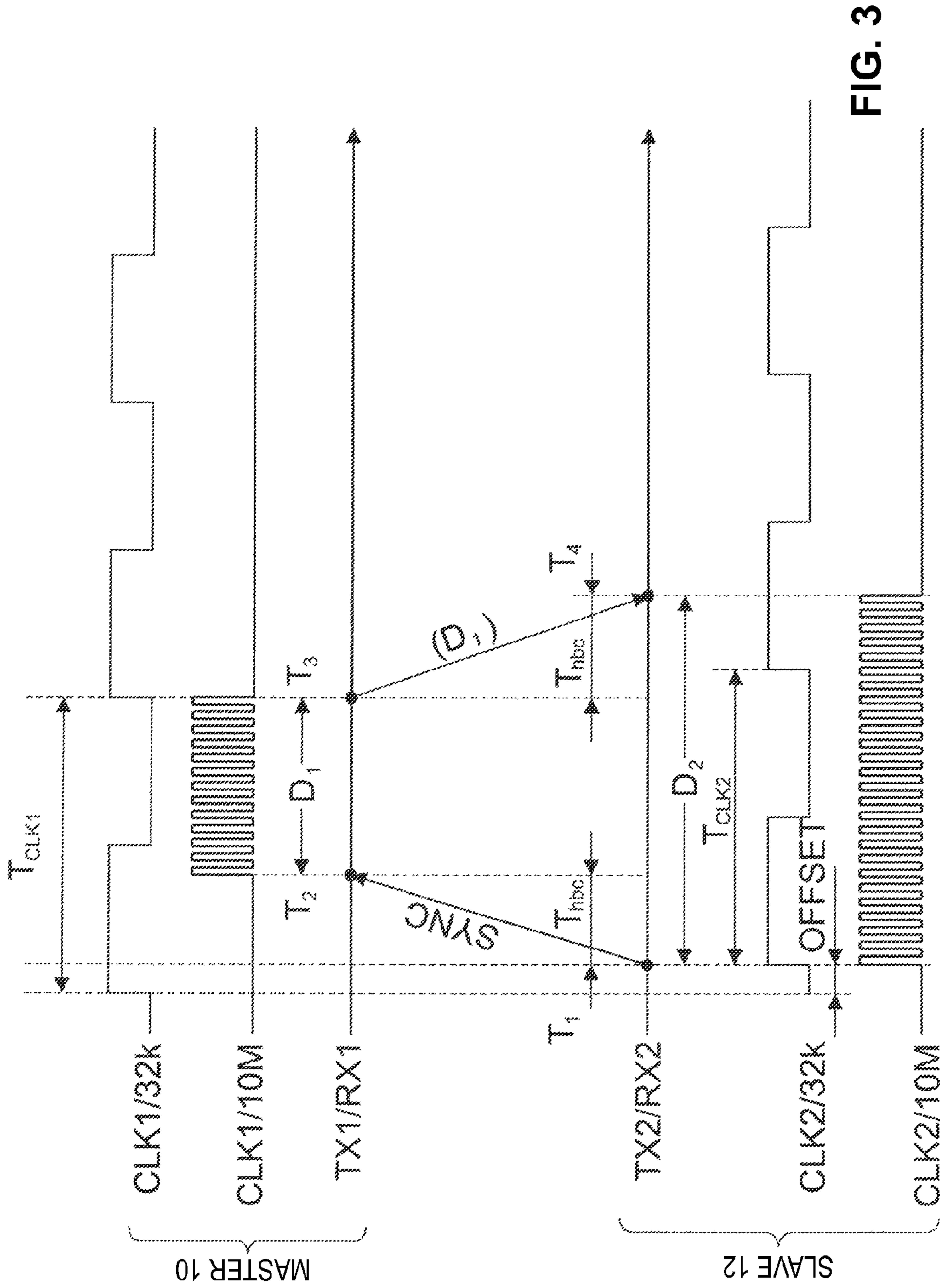


FIG. 3

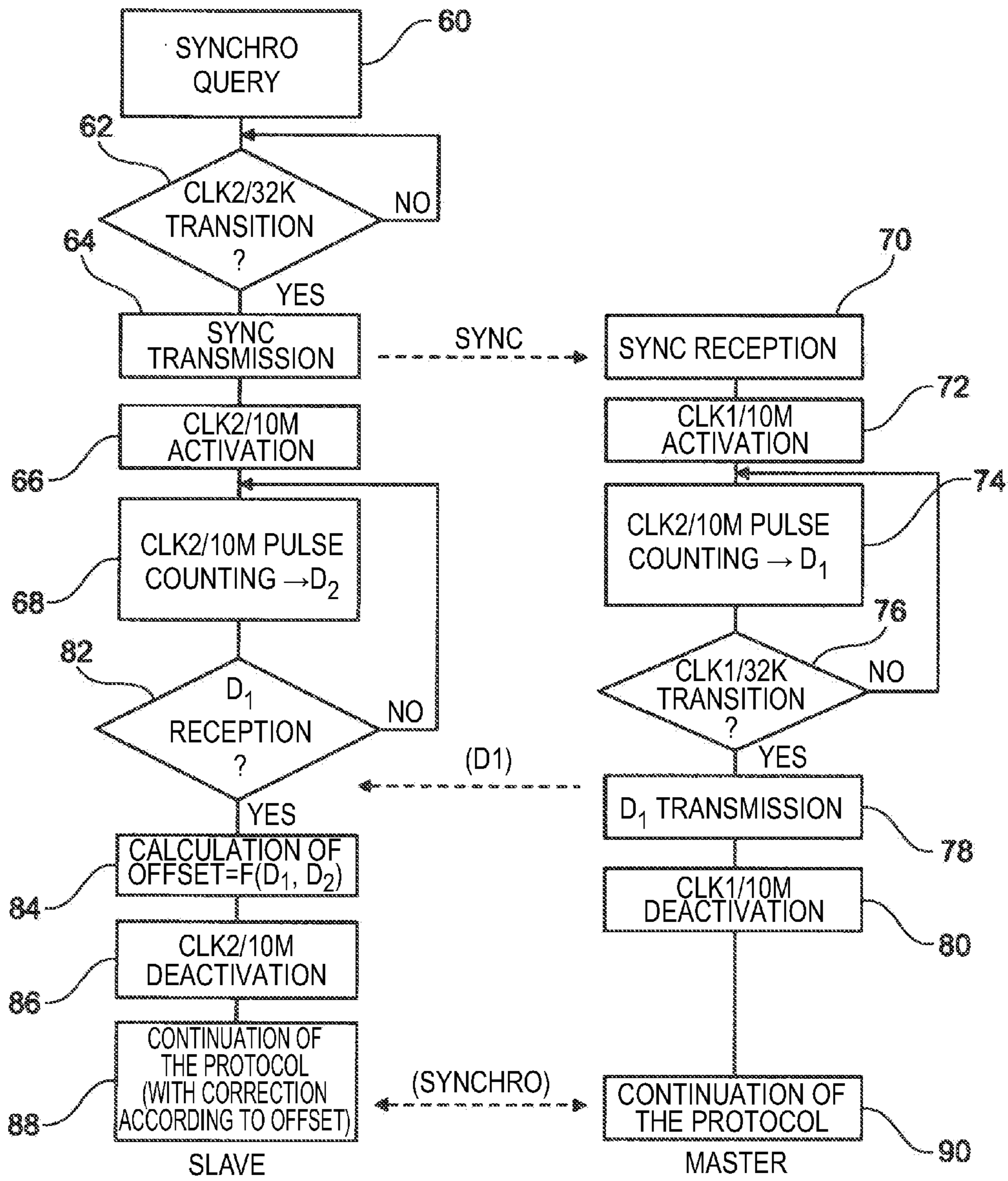


FIG. 4

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**APPARATUS AND METHOD FOR
QUANTIFICATION OF THE
DESYNCHRONIZATION BETWEEN THE
CLOCKS OF TWO HBC ACTIVE IMPLANTS**

FIELD

The present invention relates to “medical devices” as defined by the 14 Jun. 1993 Directive 93/42/EEC of the Council of the European Communities, and more specifically to the “active implantable medical devices” as defined by the 20 Jun. 1990 Directive 90/385/EEC of the Council of the European Communities, such as those implanted devices that continuously monitor a patient’s cardiac rhythm and deliver if necessary to the heart electrical pulses for stimulation, cardiac resynchronization, cardioversion and/or defibrillation in case of a rhythm disorder detected by the device, as well as neurological devices, cochlear implants, devices for pH measurement, and devices for intracorporeal impedance measurement (such as the measure of the transpulmonary impedance or of the intracardiac impedance).

BACKGROUND

The present invention relates more particularly to devices that implement autonomous implanted capsules that are free of any physical connection to another medical device, which may be implanted (such as the case of a stimulation pulse generator) or not implanted (such as an external programmer or monitoring device for remote monitoring of the patient). Communication between such an autonomous implanted capsule and another medical device is typically conducted by the interstitial tissues of the body and is known as intracorporeal communication or human body communication (“HBC”).

These autonomous implanted capsules using HBC signals for communications are for this reason called “leadless capsules” to distinguish them from the electrodes or sensors placed at the distal end of a lead, the lead being traversed throughout its length by one or more conductors connecting by galvanic conduction the electrode or the sensor distal to a generator connected at the opposite, proximal end, of the lead. Such leadless capsules are for example described in U.S. 2007/0088397 A1 and WO Patent Publication No. 2007/047681 A2 (Nanostim, Inc.) or in U.S. Patent Publication No. 2006/0136004 A1 (EBR Systems, Inc.).

These leadless capsules (which for convenience also are referred to herein simply as “capsules”) can be epicardial capsules fixed to the outer wall of the heart, or endocardial capsules fixed to the inside wall of a ventricular or atrial cavity. Capsule attachment to the heart wall is usually obtained by a protruding anchoring helical screw, axially extending out of the body of the capsule and designed to penetrate the heart tissue by screwing to the implantation site.

This capsule typically includes detection/stimulation circuitry to collect depolarization potentials of the myocardium and/or to apply electrical pulses to the site where the capsule is located. In such case, the capsule includes an appropriate electrode, which can be included in an active part of the anchoring screw. It can also incorporate one or more sensors for locally measuring the value of a parameter such as the oxygen level in the blood, the endocardial cardiac pressure, the acceleration of the heart wall, the acceleration of the patient as an indicator of activity, etc.

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It should be understood however that the present invention is not limited to a particular type of capsule, and is equally applicable to any type of leadless capsule, regardless of its functional purpose.

Of course, for the remote exchange of data, the leadless capsules incorporate transmitter/receiver circuitry for wireless communication. Several techniques have been proposed for wireless communication between the autonomous implanted capsules and a main remote device to centralize the information collected by the capsule and send to the capsule, if necessary, appropriate controls (particularly where the capsule may be an implanted pacemaker, defibrillator or resynchronizer, a subcutaneous defibrillator, or a long-term event recorder).

Thus, U.S. Patent Publication No. 2006/0136004 A1 proposes to transmit data by acoustic waves propagating inside the body. Although this technique is safe and effective, it nevertheless has the disadvantage of requiring a relatively high emitting power given the attenuation of acoustic waves into the body, and therefore allows only relatively low data transmission rates.

U.S. Pat. No. 5,411,535 proposes another communication technique, based on the use of radiofrequency waves (RF). Again, a relatively high transmission power is required, and the attenuation of these waves by intracorporeal tissue is a major barrier to their spread.

Another communication technique has been proposed by U.S. Pat. No. 4,987,897, but it is a data exchange with an external device (programmer), through the patient’s skin rather than an intracorporeal transmission. This transmission is over a short distance, between, on the one hand, the housing of a pacemaker implanted in a subcutaneous pocket and, on the other hand, an external programmer placed against the skin near the generator. Currents therefore circulate through the skin in a very distant area from the sensitive areas, particularly in a very distant area from the myocardium, which avoids any risk of disruption of the natural or stimulated depolarization waves of the latter.

The U.S. Patent Publication No. 2007/0088397 A1 also proposes to use the stimulation pulses produced by a capsule as a vehicle for the transmission of data previously collected or created by the capsule. To this purpose, the pulse, instead of presenting a monotonic variation of voltage, is interrupted in a controlled manner for very short durations in order to create (modulate) in the profile of the pulse very narrow pulses whose sequence corresponds to binary encoding of information to be transmitted.

The processing of signals exchanged between the capsules and/or the other medical device (e.g., an implanted pacemaker or defibrillator) or, more generally, between the various independent devices implanted in the patient’s body or not, implies to synchronize the respective processing circuits of these devices, or at least to determine their degree of desynchronization. If it is known, such a synchronization parameter may be included in the various signal processing calculations, or serve to apply an appropriate temporal correction to the timing of the received or transmitted signals. This “software synchronization” thus can be achieved which does not need to apply a corrective action or adjustment of the clock circuits of the various devices.

Further, to the extent that the various devices are all physically independent, each has its own clock that provides control of the various circuits, including the circuits for digital signal processing. These independent clocks have a frequency difference which, even if it is small, introduces over time a desynchronization of these devices that becomes necessary to identify and compensate for.

In a related area, U.S. Patent Publication No. 2009/030484 A1 proposes to synchronize the sequencing of two separate hearing aids, arranged on both ears of a patient, to reduce mutual interference that may be generated. These hearing aids are equipped with their own clocks, and with means for exchanging resynchronization data to readjust these two clocks.

The problem of synchronous operation of leadless implants is also discussed by US Patent Publication No. 2007/0255330 A1, which refers to measuring the time interval between the detection in two different regions of the patient of the same physiological event, or by US Patent Publication No. 2005/0197680 A1, which acts on a phase-locked loop for fine control of the clock frequency.

The frequency drift is increased by the fact that, to avoid excessive consumption, the implanted devices use clocks operating at a relatively low frequency (typically in the range of 32 kHz) and thus with a lower temporal resolution (about 30 μ s) which does not allow a detailed evaluation of the synchronization timing differences. This is particularly mentioned, as such, by the US Patent Publication No. 2007/0088394 A1, but outside of the context of synchronization between the implants.

Indeed, whatever the technique used, the processing of the received or emitted HBC signal by a capsule requires considerable energy compared to the energy resources available to the capsule. Given its autonomous nature, the capsule can in fact use only its own resources such as an energy harvester circuit (by the movement of the capsule) and/or a small integrated battery, which energy sources are in themselves well known to persons of ordinary skill in the art and form no part of the present invention, and therefore are not described in detail.

The management of the available energy is, however, a crucial point in the implementation of HBC signals to and from autonomous implanted capsules, and it is essential to develop techniques that minimize the energy requirements of these capsules.

OBJECTS AND SUMMARY

It is, therefore, an object of the present invention to determine an offset between the clocks of two independent medical devices in wireless communications, so as to implement any appropriate remedial actions.

It is a further object of the present invention to address the delay of transmission of the pulses from one device to another.

Broadly, one aspect of the present invention is directed to an apparatus and method for quantifying a desynchronization between two clocks of a system comprising at least one medical device and at least one active implantable medical device of the general type disclosed in US Patent Publication No. 2007/0088397 A1 cited above, which disclosure is incorporated herein by reference, that is to say two devices that wirelessly communicate with each other via intracorporeal communications, preferably using signals consisting of modulated electrical pulse trains carried by the interstitial tissues of the body.

Although the present invention is described in the context and example of using HBC signals for wireless communication, it is not limited to this specific mode of transmission of pulses, and also applies to the case of signals consisting of other types of waves conducted by the interstitial tissues of the body, or of acoustic waves or magnetic or electromagnetic pulses or waves passing through the body.

One embodiment of such a system comprises: a first device comprising a first clock, first signal transmitter/receiver, and a first digital processing circuit that is clocked by the first clock; and a second device comprising a second clock, a second signal transmitter/receiver, and a second digital processing circuit clocked by the second clock. The term “clocked” as used herein means to control the speed at which the respective digital processing circuits operate to process digital data, make calculations, execute instructions, etc., that is a function of the clock pulse period or the corresponding clock pulse rate of the first or second clocks. In this embodiment, one of the first and second devices also comprises a third clock, the third clock having a higher frequency and thus a shorter period and a faster clock pulse rate (or speed) than the first and second clocks, and the first and second clocks have essentially the same period or clock pulse rate. To quantify at the second device a lack of synchronism of the second clock as compared to the first clock, the third clock is used in the following manner:

- a) Using the third clock of one of the first and second devices, on a predetermined transition of the second clock, transmitting from the second device to the first device a synchronization query signal;
- b) counting the pulses of the third clock that occur from the predetermined transition of the second clock initiating the synchronization query signal until the detection of a predetermined transition of the first clock; then
- c) transmitting from the first device to the second device a response signal to the synchronization query; and
- d) at the second device, upon reception of the response signal to the synchronization query, calculating a temporal shift corresponding to a lack of synchronism based on the result of counting the pulses of the third clock.

In a preferred embodiment, the first device includes the third clock, and counting the pulses of the third clock further comprises:

counting the pulses of the third clock until detection of said predetermined transition of the first clock, wherein the response signal to the synchronization request contains the result based on counting the pulses of the third clock and.

In an alternate preferred embodiment, the second device includes the third clock and:

the second device counts the pulses of the third clock until the reception of said response signal to the synchronization request.

In either of these two alternative preferred embodiments, the calculation of the lack of synchronization can be achieved, regardless of the propagation time of the conducted electrical pulses between the first and second devices, by application of a relation of the form:

$$\text{OFFSET} = T_{clk} - D,$$

wherein “OFFSET” is the value of a temporal offset corresponding to a lack of synchronization, “ T_{clk} ” is the period of the first and second clocks, and D is a duration corresponding to the result from counting the pulses of the third clock.

In another alternative embodiment of the present inventions one of the first and second devices includes the third clock and the other of the first and second device includes a fourth clock, the fourth clock having the same clock frequency and pulse rate as the third clock, wherein:

Step a) further comprises starting the counting of the pulses of the fourth clock on the predetermined transition of the second clock;

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Step b) further comprises the counting the pulses of the third clock until detection of said predetermined transition of the first clock;

The response signal to the synchronization request emitted in step c) contains the result of the counting of the pulses of the third clock; and

Step d) includes the calculation of said temporal delay based on the results of both the counting of the pulses of the fourth clock and of the counting of the pulses of the third clock.

In the latter case, the calculation of the desynchronization can be achieved, taking into account the propagation time of the electrical pulses conducted between the first and second devices, by applying a relation of the form:

$$\text{OFFSET} = T_{clk} - [(D_1 + D_2)/2],$$

wherein "OFFSET" is the offset value corresponding to said lack of time synchronization, " T_{clk} " is the period of the first and second clocks, D_1 is the duration corresponding to the result of counting of the pulses of the third clock, and D_2 is the duration corresponding to the result of counting of the pulses of the fourth clock.

Advantageously, in a preferred embodiment of the various embodiments described above, the third and/or fourth clocks are selectively activated and initially deactivated clocks, while the first and second clocks are continuously activated, wherein the method:

in step a) includes a sub-step of activating of the fourth clock before the start of the counting of pulses, and in step d) includes a sub-step of disabling the fourth clock after receiving the response signal to the synchronization request; and/or

in step b) includes a sub-step of activating of the third clock before the start of the counting of pulses, and of disabling of the third clock after detection of the predetermined transition of the first clock.

The frequencies of the first and second clocks defining the corresponding periods are preferably equal and second to be between 15 and 50 kHz, more preferably between 30 and 35 kHz, and the fast clock frequencies of the third and fourth clocks are preferably at least 10 times the frequency of the first and second slow clocks. The accuracy of the slow clocks is advantageously better than 1%.

As noted above, the transmission of information takes place by propagation of pulses in the interstitial tissues of the patient's body, and this propagation requires a significant delay. Accordingly, a propagation delay also could be added to the required response time of the low noise amplifier ("LNA") conventionally found in an HBC signal receiver circuit to detect a pulse at the input of the receiver circuit.

These two added delays lead to a "transmission time" that most often cannot be ignored, and which is in addition to the proper synchronization of the clocks.

It should be understood that while the result may be a count of the number of fast pulses occurring during the defined period of time, in the alternative the result may be a value corresponding to the counted number of pulses, or it may be a value corresponding to a duration of time that is based on the number of pulses counted during the defined period. The precise data constituting the result transmitted is a matter of design choice based on the signals being used to transmit the result from one device to another.

Another aspect of the present invention is directed to a medical device that can be used in a system of the type described above involving at least two medical devices, such that the one medical device has the capability to communicate

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wirelessly with system to determine a quantification of its desynchronization with the system with which its communicates.

One such embodiment is directed to an active implantable medical device able to communicate wirelessly by pulse trains conducted by interstitial tissues of a patient (HBC signals) with at least one other medical device, including apparatus for quantifying a desynchronization of said active implantable medical device from said other medical device, comprising:

a transmitter/receiver of HBC signals;

a slow clock having a slow clock rate signal having pulses including transitions;

a first digital signal processing circuit;

means responsive to a predetermined transition of said slow clock for transmitting a first HBC signal that is a synchronization query signal (SYNC);

means for receiving a second HBC signal that is a response to a SYNC signal, including a first count of pulses of a fast clock rate signal occurring during a period corresponding to a period of said slow clock rate signal, the fast clock rate signal have a faster pulse clock rate than said slow clock; and

means for computing a temporal shift (OFFSET) corresponding to said desynchronization based on said first count.

In a preferred embodiment, the means for computing the temporal shift based on said first count applies the relation of the form:

$$T_{clk} = \text{OFFSET} - D_1$$

wherein OFFSET is a temporal shift value corresponding to said desynchronization, T_{clk} is a period of the slow clock rate signal and D_1 is a duration corresponding to the count of the fast clock rate signal pulses.

Preferably, the active implantable medical device further comprises:

a second clock having said fast clock pulse rate signal having pulses and transitions;

means for determining a second count corresponding to a number of pulses of said second clock fast clock pulse rate signal occurring during a period between the transmission of the first HBC signal and the receipt of the second HBC signal;

wherein the means for computing the temporal shift further comprises means for computing said temporal shift based on said first count and said second count.

Further, in this embodiment the means for computing the temporal shift applies the relation of the form:

$$\text{OFFSET} = T_{clk} - [(D_1 + D_2)/2],$$

wherein OFFSET is a value shift value corresponding to said desynchronization, T_{clk} is the period between the slow clock rate signal pulses, D_1 is the duration corresponding to the first count and D_2 is the duration corresponding to the second count.

In one embodiment, the active implantable medical device is a capsule wherein said second clock has an active state and an inactive state further comprising means for placing said second clock in an active state in response to a transmitted first HBC signal, and for placing said second clock in an inactive state in response to a receipt of said second HBC signal.

Yet another aspect of the present invention is directed to a medical device that can be used in a system of the type described above involving at least two medical devices, such that the one medical device has the capability to communicate

wirelessly with system and provides data to the system to permit the other medical devices in the systems to determine their quantification of desynchronization with the system. One such embodiment is directed to a medical device able to communicate wirelessly by pulse transition conducted by interstitial tissues of a patient (HBC signals) with at least one active implantable medical device, an apparatus for use in quantifying desynchronization of said active implantable medical device from said medical device comprising:

- a transmitter/receiver of HBC signals;
- a slow clock having a slow clock pulse rate signal having pulses and transitions;
- a fast clock having a fast clock pulse rate signal having pulses and transitions, said fast clock pulse rate being faster than the slow clock rate;
- means responsive to a receipt of a first HBC signal corresponding to a synchronization query (SYNC) from said active implantable medical device for determining a first count corresponding to a number of fast clock rate signal pulses counted during a period between two consecutive pulses of said slow clock pulse rate signal;
- means responsive to determining said first count for transmitting a second HBC signal corresponding to a response to said first HBC signal and including said first count.

In a preferred embodiment, the medical device is an active implantable medical device such as a cardiac pacemaker, or defibrillator or a long term recording device.

DRAWINGS

Further features, characteristics and advantages of the present invention will become apparent to a person of ordinary skill in the art from the following detailed description of preferred embodiments of the present invention, made with reference to the drawings annexed, in which like reference characters refer to like elements, and in which:

FIG. 1 is a schematic illustration of a system of medical devices including leadless capsules implanted within the body of a patient;

FIG. 2 is a functional block diagram showing the different constituting stages of two devices of the system of FIG. 1 and their communication method;

FIG. 3 illustrates a series of timing diagrams representative of a preferred operation for quantifying a desynchronization of the two devices; and

FIG. 4 illustrates, in parallel, two flow chart diagrams corresponding to different steps respectively implemented in each of the two medical devices of FIG. 2 for implementing a preferred embodiment of the present invention.

DETAILED DESCRIPTION

With reference to the drawings, an example of a system and method, and medical devices that are part of such a system, according to preferred embodiments of the present invention will now be described.

With reference to FIG. 1, a system of medical devices implanted in the body of a patient is shown, communicating with each other through wireless communication of signals through the intracorporeal tissues via human body communication (“HBC”).

Implanted in the patient are, for example, an implantable device **10** such as an implantable defibrillator/pacemaker/resynchronizer or a subcutaneous defibrillator or a long-term event recorder. In this embodiment, device **10** is part of a system comprising a plurality of devices **12** to **18**, with which

device **10** is likely to communicate by HBC. It should be understood that device **10** can also be implanted as a capsule, without requiring the presence of a subcutaneously implanted defibrillator/pacemaker/resynchronizer connected to one or more leads, or a subcutaneous defibrillator.

The devices **12** to **18** may include intracardiac capsules **12** and/or epicardial capsules **14**, located directly on the patient’s heart. Such capsules are for example described in US Patent Publication No. 2007/0088397 A1, WO 2007/047681 A2 and US Patent Publication No. 2006/0136004 A1, and are attached to the heart wall by a protruding anchoring screw intended to penetrate into the cardiac tissue by screwing at the implantation site. The anchoring screw may be a passive screw, serving only to fix the capsule, or an active screw, serving to fix the capsule as well as to collect the signals of depolarization propagated in myocardium tissues and/or deliver electrical pulses to the implantation site.

The system may also include other devices such as myopotential sensors **16** or neurological stimulation devices, and possibly an external device (**18**) disposed on an armband and provided with electrodes in contact with skin. The device **10** can be used as a gateway to the outside world to communicate with an external device **20** such as a programmer device or a remote data transmission device with which it can communicate, notably by RF telemetry in the Medical Implants Communication System (MICS) frequency band 402-405 MHz, or in the 401-402 MHz and 405-406 MHz MEDS frequency bands, or in the public ISM (Industrial, Scientific and Medical) 863-870 MHz, 902-928 MHz and 2.4 GHz frequency bands used by medical devices.

Each of the devices **10** to **18** is provided with at least one pair of electrodes which for the implantable devices are in direct contact with body tissues, or in the case of an external device **18** in contact with the skin or connected to leads or having electrodes on the connector or on the housing for the device **10**.

One of the devices, for example, the pacemaker/defibrillator **10** may act as a hub or master device in a star wireless network architecture, in which the various leadless capsules (**12**) to (**18**) are slave devices—this for a communication from the pacemaker/defibrillator to the capsule; for a communication in the other direction, the roles are reversed.

The nature of the communication between the different devices, networked or not, and the type of network considered, however, do not form any part of the present invention, which instead concerns only the initial communication between any of the two implanted devices.

The following example of communication between the pacemaker/defibrillator **10** as a master device and an intracardiac capsule **12** as a slave device is provided using HBC signals, but this example is purely illustrative and non limiting and it should be understood that the present invention can be applied to any wireless communication between any two medical devices (**10**) to (**20**) described above, and this application is indifferently as to the direction of the communication (that is to say that the roles of master and slave can be reversed).

Referring to FIG. 2, a schematic of the internal circuit of the pacemaker/defibrillator **10** and of the leadless capsule **12** required for the implementation of the present invention is illustrated. In the following designations, only the constituting components **22** to **36** of the leadless capsule **12** are described, and it should be understood that the pacemaker/defibrillator **10** includes similar elements **42** to **56**, corresponding to the elements **22** to **36** described in the case of the capsule, that work and function in the same way.

Capsule **12** includes two electrodes **22**, **24** in direct contact with body tissues, allowing the transmission and reception of HBC signals consisting of pulses carried by the interstitial tissues of the body located between the two devices **10** and **12** and also used for detection and pacing. For the capsule **12**, one of the electrodes **22** or **24** may be constituted by the anchoring screw into the tissue of the heart, or by one of the electrodes on the housing or connected to the housing of the capsule. The pair of electrodes **22**, **24** is connected to a receiver circuit (RX) **26** having for example a low noise amplifier (not shown) for detection of the potentials collected between the electrodes and to a transmitter circuit (TX) **28** to produce pulses for communication by HBC signals with the remote device (**10**). Transmitter (TX) and receiver (RX) circuits are conventional and well known to a person of ordinary skill in the art, and may be combined into an integrated circuit structure. For example, reference here is made to the following publications: as to the design of a receiver “CMOS RF Receiver Design for Wireless LAN Applications”: <http://www.ee.ucla.edu/~brweb/papers/Conferences/RRAWCON99.pdf>; as to the design of an RF transmitter, “RF and Microwave Transmitter Design”: <http://www.wiley-vch.de/publish/en/books/forthcomingTitles/EE00/0-470-52099-Xf>; “Design data for radio transmitters and receivers”: http://books.google.de/books?id=ZsE4m6AAY7UC&redir_esc=y; and as to the design of RF transceivers which incorporate a transmitter and receiver into a single integrated circuit structure; “Circuit design for RF transceivers”; http://books.google.de/books/about/Circuit_design_for_RF_transceivers.html?id=0pGBG2E4MSgC&redir_esc=y.

The electrodes **22**, **24** can also be connected to an electrical pulse generator circuit (not shown) if the capsule **12** incorporates this function and/or to a detection circuit for collection of spontaneous or stimulated cardiac depolarization potentials (these latter potentials are to be distinguished from the potentials corresponding to the HBC signals, the latter of which are emitted at an energy level far below that of cardiac potentials and that of myopotentials of the body). In other words, the electrodes **22**, **24** can perform several functions, namely the collection of cardiac potential and/or the stimulation (if applicable), and in any case the transmission/reception for the HBC communication. These detection and collection circuits are well known by those skilled in the art and not involved in the present invention, and therefore have not been illustrated in FIG. **2**.

The capsule can also be provided with a sensor such as an acceleration sensor, a pressure sensor, a hemodynamic sensor, a temperature sensor, an oxygen saturation sensor, etc. It is preferably powered by an energy source such as an energy harvester circuit, a small battery, or a combination thereof.

In accordance with one embodiment of the present invention, typically, the device **12** has two separate clocks, namely a slow clock **30** and a fast clock **32**. The terms “fast” and “slow” should be understood in their relative sense, that is to say that the clock frequency of fast clock **32** is higher (and the pulse period is shorter) than that of the slow clock **30**, preferably with a difference of at least an order of magnitude (ten times).

The slow clock **30** produces pulses CL**23** at a frequency of, for example, 32,768 Hz, while the fast clock **32** produces pulses CL**21** at a frequency of 9.83040 MHz, that is to say three hundred times that of the slow clock **30**. The precise values of these frequencies are given only as non-limiting examples, and were chosen to provide frequencies having a simple, easy to be generated ratio by frequency division. For simplicity of presentation, it is simply referred to “32 kHz”

and “10 MHz” frequencies, the precise value chosen being not in itself important for the implementation of the invention. Furthermore, the pulses corresponding to these frequencies of 32 kHz and 10 MHz generated within the device **12** are designated CLK**2/32k** and CLK**2/10M**, respectively; those generated within the device **10** are designated CLK**1/32k** and CLK**1/10M**, respectively.

The slow and fast clock signals are generated in the form of square wave pulses with a 50% duty cycle, as shown in the timing diagrams in FIG. **3**. The corresponding periods are of the order of 30 μ s for the slow clock and 0.1 μ s for the fast clock.

The slow clock **30** continuously generates pulses, and these pulses are used for the sequencing of a central processing unit, such as a microprocessor or microcontroller (not shown) including all the electronics, memory, registers, etc. for controlling the various functions of the capsule, storing the collected signals, etc. The slow clock **30** thus generates the required clock signals including (i) the analysis of depolarization signals collected by the electrodes, to the generation of the electrical pulses, etc., and (ii) to communication with other implanted devices, including the master device constituted by the implanted pacemaker/defibrillator device **10** in this embodiment.

The fast clock **32**, in contrast, is only temporarily activated under the conditions which are described below. The normal state of fast clock **32** is a disabled state, and its activation is controlled by a circuit **34** controlled from data D_1 received from the master device **10** and from data D_2 developed internally by the slave device **12**, including a circuit counter **36** that counts the pulses CLK**2/10M** of the fast clock **32**. Circuit **36** is controlled by (i) the slow clock **30** and by (ii) HBC signals received from the master device **10**.

A relatively low frequency is chosen for clock **30** so as to limit consumption and manage the energy of the devices. For this reason, the fast clock **32** is enabled only when necessary, as its energy consumption is much higher and would be incompatible with its continuous operation, because the energy resources of the device are very limited (in particular, using an energy harvester circuit for the leadless capsule, or a very long life battery (10 years) for the pacemaker/defibrillator **10**).

The master device (here the pacemaker/defibrillator) **10** includes the same elements **42** to **56** as the elements (**22**) to (**36**) of the slave device constituted by the leadless capsule **12**, as mentioned above.

Thus, each of the two representative devices **10** and **12** in this exemplary system is provided with its own slow clock **30** or **50**, respectively, which ensures the sequencing of various digital circuits of each of the devices. These two clocks are physically independent, and hence they have some degree of desynchronization or a temporal offset. This offset must be calculated in order to take it into account later in the protocol for the desired operation of the system, in particular so that the windows of reception and transmission of wireless communication between master and slave devices coincide as closely as possible.

The determination of this temporal offset in the slave device **12** will now be described in detail, with reference to FIGS. **3** and **4**. On the timing diagrams of FIG. **3**, the temporal offset is indicated by the parameter OFFSET (to be determined) and is present between the signals CLK**1/32k** and CLK**2/32k** of the slow clocks.

The slave device **12** which has to be synchronized to the master device **10** waits for a predetermined transition (e.g., a rising edge) of the slow clock signal CLK**2/32k** to deliver, at

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time T_1 , a synchronization query pulse SYNC towards the master device **10** (steps **60**, **62**, **64** on the flow chart in FIG. 4).

The master device **10** receives this synchronization query pulse SYNC (step **70**) at time T_2 , with, compared to the moment of transmission, a delay $(T_2 - T_1) = T_{hbc}$ corresponding to the transmission time of the pulse through the interstitial tissues of the body from the slave device **12** to the master device **10**.

Upon reception of the pulse SYNC, the master device **10** activates its fast clock **52** (step **72**), triggered by the control circuit **54** after detection of the SYNC pulse by the receiver circuit **46**. The master device **10** then counts the number of pulses CLK1/10M generated by fast clock **52** by the counting circuit **56** (step **74**).

Meanwhile, from the delivery of the pulse SYNC, the slave device **12** activates its fast clock **32** (step **66**) and begins to count the pulses CLK2/10M of the fast clock in its counter circuit **36** (step **68**).

At master device **10**, the counting of pulses CLK1/10M of the fast clock continues until detection, at time T_3 , of a predetermined transition (e.g. a rising edge) of the signal CLK1/32k of the slow clock **50**. The number of pulses counted gives a numerical value of data D_1 representative of the duration between the times T_2 and T_3 . With a fast clock signal CLK1/32k clocked at 10 MHz, the correspondence is of 0.1 μ s per counted clock pulse.

This data value D_1 is then sent to the slave device **12** in a response message (step **78**). Subsequently, the fast clock **52** is turned off (step **80**) to limit the energy consumption of the device. The slow clock **50**, however, continues to produce the necessary pulses CLK1/32k for controlling the various digital circuits of the master device **10**, in particular for the continuation of the protocol for exchanging data with the slave device **12** as well as its other functions.

At the slave device **12**, the response message received back and containing the value D_1 is received at time T_4 , shifted from T_3 by a duration corresponding to the transmission time T_{hbc} from the master device **10** to the slave device **12**, with a delay $(T_4 - T_3) = T_{hbc}$ (assuming that the transmission time is the same in one direction or the other).

Note that the detection circuit **26** of the slave devices is initiated only during the reception window triggered by the fast clock signal CLK2/10M of the slave device. The duration of this window is a number of periods greater than or equal to one of the periods of signal CLK2/10M.

Upon reception of the returned response message (step **82**), the slave device calculates the OFFSET value from the value received D_1 and from $D_2 = (T_4 - T_1)$ internally calculated and corresponding to the number of pulses CLK2/10M of the slow clock **30** counted by the counter circuit **36** (step **84**).

The fast clock **32** is then deactivated (step **86**) to limit the energy consumption and the signal exchange protocol can then be continued (step **88**) to exchange signals with the master device **10** (step **90**). This communication can be established using the OFFSET parameter and then applying a temporal correction corresponding to the dating of the signals received or transmitted (e.g., a software synchronization).

The offset value OFFSET is calculated as follows. Assuming that the transmission time T_{hbc} is identical in both directions of communication, then:

$$T_4 - T_1 = 2 \times T_{hbc} + D_1 = D_2$$

D_2 and D_1 being known (D_2 internally counted by the slave device **12** and D_1 remotely counted by the master device **10** and transmitted to the slave device (**12**)), and it can be deduced:

$$T_{hbc} = (D_2 - D_1) / 2.$$

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The crystal oscillators used in these fast and slow clocks to set the operating frequencies have a typical accuracy of 50 ppm, for a frequency of 32,768 Hz. This accuracy is sufficient, on the scale of the transmission of a data packet via HBC signals, which can tolerate a lack of synchronization of a fast clock period due to the tolerance of 50 ppm of quartz. Thus, if T_{clk1} and T_{clk2} respectively denote the periods of the slow clocks **30** and **50** of the slave **12** and master **10** devices, it can be considered that $T_{clk1} = T_{clk2} = T_{clk}$. Thus:

$$T_3 - T_1 = T_{clk} - \text{OFFSET} = D_1 + T_{hbc},$$

hence:

$$\text{OFFSET} = T_{clk} - D_1 - T_{hbc} = T_{clk} - (D_1 + D_2) / 2$$

The OFFSET shift having thus been calculated, the window of communication with the master device can be determined and adjusted to obtain the desired timing for the transmission of data packet.

Taking into account the propagation time, T_{hbc} is optional, and depends on the desired accuracy. This delay can very roughly be estimated at about 500 ns, so that if the fast clock operates at a frequency of 1 MHz instead of 10 MHz, this delay can be neglected since it has virtually no impact on the calculation result of the OFFSET shift.

In this case, it may be considered that the moments T_1 and T_2 are mingled, as well as the moments T_3 and T_4 , and thus $D_1 = D_2$. As a result, it is no longer necessary to measure D_2 (in which case step **68** can be ignored), and directly:

$$\text{OFFSET} = T_{clk} - D_1$$

To the extent that there is no need to measure D_2 , slave device **12** may be devoid of a fast clock, which simplifies the design, reduces the size and reduces the energy consumption of slave devices.

Alternatively, since $D_1 = D_2$, if the slave device **12** is provided with a fast clock, D_1 can be evaluated using this fast clock, which gives the value of D_2 , equal to D_1 . Even if this solution requires the presence of a fast clock in slave device **12**, it simplifies the protocol for exchanging signals with master device **10**: indeed, slave device **12** has just to detect the receipt of a marker from master device **10** corresponding to the detection of the transition of the slow clock of the latter, without the need to transmit from the latter a digitized value (i.e., the value of D_1).

Generally, master device **10** may simultaneously resynchronize a set of slave devices in a system of such devices by delivering a synchronization pulse on a rising edge of CLK1/32K that is detected by each of the slave devices **14** or **12**, for example. Each slave device n starts counting pulses CLK n /10M at the detection of the synchronization pulse SYNC from a relative master device **10**, until the next rising edge of pulse CLK n /32K.

Master device thus **10** preferably maintains slave devices **12** or **14** regularly synchronized by delivering a synchronization pulse SYNC, for example, every 2 ms.

Advantageously, the slave device **12** or **14** may use such a synchronization pulse SYNC as an internal clock. In such a case, slave device **12** or **14** does not need a crystal oscillator as an internal clock. Hence a gain in energy consumption and size results. When the slave device needs to perform particular operations at a faster speed, it can set off (i.e., turn on) a fast resistor-capacitor (RC) circuit having an output pulse signal that acts as a fast clock that it interrupts (deactivates) upon completion of the particular higher speed operation processing.

Advantageously, in one embodiment, the stimulation and the discharge of stimulation capacitors of the master and slave

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devices can be synchronized to the 2 ms clock of the master device, allowing them to serve as synchronization pulses.

One skilled in the art will appreciate that the foregoing embodiments are provided for purposes of illustration and not of limitation, and that variations in the specific values of parameters provided may be made by persons of ordinary skill in the art without departing from the nature and scope of the present invention.

The invention claimed is:

1. A method for quantification of a desynchronization between two clocks of a system comprising at least two active implantable medical devices that communicate wirelessly, including:

a first active implantable medical device comprising a first slow clock having a slow clock pulse rate, a first transmitter/receiver, and a first digital processing circuit clocked by the first slow clock; and

a second active implantable medical device comprising a second slow clock having said slow clock pulse rate, a second transmitter/receiver, and a second digital processing circuit clocked by the second slow clock, wherein the first active implantable medical device and the second active implantable medical device are configured to communicate wirelessly by human body communication signals (HBC signals), wherein the HBC signals comprise pulse trains conducted by interstitial tissues of the body, and wherein the first transmitter/receiver and the second transmitter/receiver are configured to transmit and receive the HBC signals;

wherein the first active implantable medical device includes a first fast clock having a fast clock pulse rate, the fast clock pulse rate being faster than the slow clock pulse rate;

the quantification method comprising the steps of:

a) at the second active implantable medical device, detecting a predetermined transition of a pulse of the second slow clock, and in response thereto transmitting from the second active implantable medical device to the first active implantable medical device a synchronization query;

b) at the first active implantable medical device, detecting said synchronization query and a predetermined transition of a pulse of said first slow clock, and counting a first number of pulses of the first fast clock occurring between said detected synchronization query and said detected predetermined transition of the pulse of the first slow clock; then

c) in response to counting said first number of pulses, transmitting from the first active implantable medical device to the second active implantable medical device a response signal to the synchronization query, wherein the response signal includes a result corresponding to the counted first number; and

d) receiving at the second active implantable medical device, the response signal to the synchronization query, and computing a temporal shift (OFFSET) corresponding to a desynchronization based on said counted first number of fast clock pulses;

wherein step b) further comprises selectively activating the first fast clock before the start of counting of the first fast clock pulses, and deactivating the first fast clock after detection of the predetermined transition of the pulse of the first slow clock.

2. The method of claim 1, wherein said quantification of desynchronization further comprises applying, regardless of

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a propagation delay (T_{hbc}) of the HBC signals between the first and second active implantable medical devices, a relation of the form:

$$T_{clk} = \text{OFFSET} - D_1$$

wherein OFFSET is a temporal shift value corresponding to said desynchronization, T_{clk} is a period of the slow clock rate signal, and D_1 is a duration corresponding to the counted first number of fast clock pulses.

3. The method of claim 2, wherein:

step a) further comprises selectively activating the second fast clock before the start of counting of the second fast clock pulses, and step d) further comprise deactivating the second fast clock after reception of the response signal to the synchronization query.

4. The method of claim 1, the second active implantable medical device having a second fast clock having a second fast clock pulse rate, wherein:

step a) further comprises counting, at the second active implantable medical device, a second number of pulses of the second fast clock occurring during a period between said predetermined transition of said pulse of said second slow clock and reception in step d) of said response signal to the synchronization query.

5. The method of claim 4, wherein said quantification of desynchronization further comprises applying, regardless of a propagation delay (T_{hbc}) of the HBC signals between the first and second active implantable medical devices, a relation of the form:

$$T_{clk} = \text{OFFSET} - D_1,$$

wherein OFFSET is a temporal shift value corresponding to said desynchronization, T_{clk} is a period of the slow clock rate signal, and D is a duration corresponding to one of the counted first and second numbers of fast clock pulses.

6. The method of claim 1, the second active implantable medical device having a second fast clock having said fast clock pulse rate, wherein:

step a) further comprises, at the second active implantable medical device, in response to the predetermined transition of the pulse of the second slow clock counting a second number of pulses of the second fast clock occurring during a period between said predetermined transition of said pulse of said second slow clock and reception in step d) of said response signal to the synchronization query;

step b) further comprises counting said first number as the number of pulses of the first fast clock occurring from said predetermined transition of the pulse of the second slow clock until detection of said predetermined transition of the pulse of the first slow clock;

the response signal to the synchronization query transmitted in step c) contains a result (D_1) corresponding to said counted first number of the first fast clock pulses; and

step d) further comprises computing said temporal shift (OFFSET) based on said counted first number and second number of fast clock pulses.

7. The method of claim 6, wherein the quantification of said desynchronization further comprises determining a propagation time of the HBC signals between the first and second active implantable medical devices, and applying a relation of the form:

$$\text{OFFSET} = T_{clk} - [(D_1 + D_2)/2],$$

wherein OFFSET is a temporal shift value corresponding to said desynchronization, T_{clk} is the period between the slow clock rate signal pulses, D_1 is a duration corre-

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sponding to the counted first number of the fast clock pulses, and D_2 is a duration corresponding to the counted second number of the fast clock pulses.

8. The method of claim 1, wherein the frequencies of the first and second slow clocks are equal and selected from between 15 and 50 kHz.

9. The method of claim 1, wherein the first fast clock rate signal is at least 10 times faster than the slow clock rate signal.

10. The method of claim 1, wherein the accuracy of the slow clocks is better than 1%.

11. In an active implantable medical device able to communicate wirelessly with at least one other medical device, an apparatus for quantifying a desynchronization of said active implantable medical device from said other medical device, comprising:

a transmitter/receiver of human body communication signals (HBC signals), wherein the HBC signals comprise pulse trains conducted by interstitial tissues of the body, and wherein the active implantable medical device is configured to communicate wirelessly with the at least one other medical device by the HBC signals;

a slow clock having a slow clock rate signal having pulses including transitions;

a first digital signal processing circuit clocked by said slow clock;

means responsive to a predetermined transition of a pulse of said slow clock for transmitting a first HBC signal corresponding to a synchronization query;

means for receiving a second HBC signal corresponding to a response to said synchronization query, including a first result corresponding to a first count of pulses of a fast clock rate signal occurring during a period corresponding to a period of said slow clock rate signal, the fast clock rate signal have a faster clock pulse rate than said slow clock;

means for computing a temporal shift (OFFSET) corresponding to said desynchronization based on said first result;

a second clock having said fast clock pulse rate signal having pulses and transitions; and

means for counting a second number of pulses of said second clock occurring during a period between the transmission of the first HBC signal and the receipt of the second HBC signal;

wherein the means for computing the temporal shift further comprises means for computing said temporal shift based on said first result and said counted second number; and

wherein said second clock has an active state and an inactive state, and wherein the apparatus further comprises means for placing said second clock in an active state in response to a transmitted first HBC signal, and for placing said second clock in an inactive state in response to a receipt of said second HBC signal.

12. The apparatus of claim 11, wherein said means for computing the temporal shift based on said first result applies the relation of the form:

$$T_{clk} = \text{OFFSET} - D_1$$

wherein OFFSET is a temporal shift value corresponding to said desynchronization, T_{clk} is a period of the slow clock rate signal and D_1 is a duration corresponding to the first result.

13. The apparatus of claim 11, wherein said means for computing the temporal shift applies the relation of the form:

$$\text{OFFSET} = T_{clk} - [(D_1 + D_2)/2],$$

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wherein OFFSET is a temporal shift value corresponding to said desynchronization, T_{clk} is the period between the slow clock rate signal pulses, D_1 is a duration corresponding to the first result and D_2 is a duration corresponding to the counted second number.

14. The apparatus of claim 11, wherein said fast clock rate signal is at least 10 times faster than said slow clock rate.

15. The apparatus of claim 11, wherein the slow clock is selected from the range of 15 and 50 kHz.

16. The apparatus of claim 11, wherein said active implantable medical device is an autonomous leadless capsule.

17. The apparatus of claim 11, wherein said active implantable medical device is an autonomous leadless capsule and said second clock further comprises a resistor capacitor circuit having said fast clock pulse rate.

18. In a first active implantable medical device able to communicate wirelessly with at least one second active implantable medical device, an apparatus for use in quantifying desynchronization of said second active implantable medical device from said first active implantable medical device, comprising:

a transmitter/receiver of human body communication signals (HBC signals), wherein the HBC signals comprise pulse trains conducted by interstitial tissues of the body, and wherein the first active implantable medical device is configured to communicate wirelessly with the at least one second active implantable medical device by the HBC signals;

a slow clock having a slow clock pulse rate having pulses and transitions;

a fast clock having a fast clock pulse rate having pulses and transitions, said fast clock pulse rate being faster than the slow clock pulse rate, and said fast clock having an active state and an inactive state;

means responsive to a receipt of a first HBC signal corresponding to a synchronization query from said second active implantable medical device for counting a first number of fast clock rate signal pulses occurring during a period between two consecutive pulses of said slow clock pulse rate signal;

means responsive to said counted first number for transmitting a second HBC signal corresponding to a response to said first HBC signal and including therein a first result corresponding to said counted first number;

means for placing said fast clock in the active state in response to a transmitted first HBC signal, and for placing said fast clock in the inactive state in response to a receipt of said second HBC signal;

means responsive to a predetermined transition of a pulse of said slow clock for transmitting a third HBC signal corresponding to a synchronization query;

means for receiving a fourth HBC signal that is a response to said third HBC signal, including a second result corresponding to a counted second number of pulses of a fast clock rate signal occurring between two consecutive pulses of said slow clock rate; and

means for computing a temporal shift (OFFSET) corresponding to said desynchronization based on said second result;

wherein said means for computing the temporal shift applies the relation of the form:

$$\text{OFFSET} = T_{clk} - [(D_1 + D_2)/2];$$

wherein OFFSET is a temporal shift value corresponding to said desynchronization, T_{clk} is the period between the slow clock rate signal pulses, D_1 is a duration corre-

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sponding to the second result and D2 is a duration corresponding to the first result.

19. The apparatus of claim 18, wherein said fast clock rate signal is at least 10 times faster than said slow clock rate.

20. The apparatus of claim 19, wherein said fast clock further comprises a crystal oscillator.

21. The apparatus of claim 19, wherein said fast clock further comprises a resistor-capacitor circuit.

22. The apparatus of claim 18, wherein the slow clock is selected from the range of 15 and 50 kHz.

23. A method for quantification of a desynchronization between two clocks of a system comprising at least two medical devices, including:

a first medical device comprising a first clock having a slow clock pulse rate signal, a first transmitter/receiver of wireless signals, and a first digital processing circuit clocked by said first clock; and

a second medical device comprising an active implantable medical device having a second clock, said second clock having said slow clock pulse rate signal, a second transmitter/receiver of wireless signals, and a second digital processing circuit clocked by the second slow clock, wherein the first medical device and the second medical device are configured to communicate with one another wirelessly by signals conducted through the tissues of a patient's body;

wherein the first medical device includes a third clock having a fast clock pulse rate signal and the second medical device includes a fourth clock having the fast clock pulse rate signal, a rate of the fast clock pulse rate signal being faster than a rate of the slow clock pulse rate signal;

the quantification method comprising the steps of:

a) at the second medical device, detecting a predetermined transition of a second clock pulse, and wirelessly transmitting from the second medical device to the first medical device a synchronization query;

b) at the first medical device, detecting said synchronization query and a predetermined transition of a pulse of the first clock, and determining a first result corresponding to a number of pulses of the third clock occurring between said detected synchronization query and said detected predetermined transition of the first clock pulse;

c) transmitting from the first medical device to the second medical device a response signal to the synchronization query; and

d) receiving at the second medical device, the response signal to the synchronization query, and computing a temporal shift (OFFSET) corresponding to a desynchronization based on the first result;

wherein:

step a) further comprises, at the second medical device, in response to the predetermined transition of the second clock, determining a second result corre-

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sponding to a number of pulses of the fourth clock occurring during a period between said predetermined transition of said pulse of said second clock and reception in step d) of said response signal to the synchronization query;

step b) further comprises determining said first result based on counting the number of pulses of the third clock occurring from said predetermined transition of the pulse of the second clock until detection of said predetermined transition of the pulse of the first clock; step c) further comprises providing said first result in the response signal to the synchronization query; and step d) further comprises computing said temporal shift (OFFSET) based on said first and second results.

24. The method of claim 23, wherein the quantification of said desynchronization further comprises determining a propagation time of the signals conducted through the tissues of the patient's body between the first and second active implantable medical devices, and applying a relation of the form:

$$\text{OFFSET} = T_{\text{clk}} - [(D1 + D2)/2],$$

wherein OFFSET is a temporal shift value corresponding to said desynchronization, T_{clk} is the period between the slow clock rate signal pulses, D1 is a duration corresponding to the first result, and D2 is a duration corresponding to the second result.

25. The method of claim 23, wherein:

step b) further comprises selectively activating the third clock before the start of counting of the third clock pulses, and deactivating the third clock after detection of the predetermined transition of the pulse of the first clock.

26. The method of claim 25, wherein:

step a) further comprises selectively activating the fourth clock before the start of counting of the fourth clock pulses, and step d) further comprises deactivating the fourth clock after reception of the response signal to the synchronization query.

27. The method of claim 23, wherein the frequencies of the first and second clocks are equal and selected from between 15 and 50 kHz.

28. The method of claim 23, wherein the fast clock pulse rate signal is at least 10 times faster than the slow clock pulse rate signal.

29. The method of claim 23, wherein the accuracy of the first and second clocks is better than 1%.

30. The method of claim 23, further comprising providing the third clock as a resistor-capacitor circuit having said fast clock pulse rate signal.

31. The method of claim 23, further comprising providing the third clock as a crystal oscillator having said fast clock pulse rate signal.

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